



SHEET 1 OF 8

FORM PTO-1449 U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE INFORMATION DISCLOSURE STATEMENT (Use several sheets if necessary)	ATTY. DOCKET NO. 4172-85	SERIAL NO. 10/679,699
	APPLICANT Bar-Or et al.	
	FILING DATE October 2, 2003	GROUP ART 1649

U.S. PATENT DOCUMENTS

*EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB CLASS	FILING DATE IF APPROP.
GE	A1.	6,096,737	8/1/2000	Loder	514	217	
	A2.	6,475,743	11/5/2002	Bar-Or et al.	435	7.1	
	A3.	6,555,543	4/29/2003	Bar-Or et al.	514	255.02	
	A4.	6,492,179	12/10/2002	Bar-Or et al.	436	74	
	A5.	6,461,875	10/8/2002	Bar-Or et al.	436	536	
	A6.	6,090,780	7/18/2000	Prasad	514	11	
	A7.	4,771,056	9/13/1988	Rozencwaig	514	325	
	A8.	4,661,500	4/28/1987	Rozencwaig	514	325	

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		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB CLASS	TRANSLATION	
							YES	NO
GE	A9.	WO 02/11676	2/14/2002	PCT				
	A10.	NZ 033544	8/31/2001	New Zealand				
	A11.	EP 0 835 660 A1	4/15/1998	EPO				
	A12.	EP 0 214 557 A2	3/18/1987	EPO				
	A13.	EP 0 214 557 A3	3/18/1987	EPO				
	A14.	WO 01/34586	5/17/2001	PCT				
	A15.	WO 00/20454	4/13/2000	PCT				

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GE	A16.	WO 02/059604	8/1/2002	PCT				
	A17.	WO 00/20840 A1	4/13/2000	PCT				
	A18.	WO 98/40748 A1	9/17/1998	PCT				
	A19.	RU2112242C1	5/27/1998	Russian Federation			X, abstract only	
	A20.	RU2125728C1	1/27/1999	Russian Federation			X, abstract only	
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GE	A22.	Esposito et al., "The Solution Structure of the C-Terminal Segment of Tau Protein," <i>Journal of Peptide Science</i> 2000, 6:550-559
	A23.	Gamblin et al., "Tau Polymerization: Role of the Amino Terminus," <i>Biochemistry</i> 2003, 42(7):2252-2257
	A24.	Crowe et al., "The N Terminal Region of Human Tau is Present in Alzheimer's Disease Protein A68 and is Incorporated into Paired Helical Filaments," <i>American Journal of Pathology</i> 1991, 139(6):1463-1470
	A25.	Berry et al., "Inhibition of Tau Polymerization by its Carboxy-Terminal Caspase Cleavage Fragment," <i>Biochemistry</i> 2003, 42:8325-8331
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GE	A28.	Bar-Or et al., "Potential Plasma Surrogate Biomarkers for CNS Demyelinating Processes," Meeting of the 19th Congress of the European Committee for Treatment and Research in Multiple Sclerosis, Sept. 17-20, 2003 (abstract first distributed at the meeting)
	A29.	Garcia-Sierra et al., "Conformational Changes and Truncation of Tau Protein During Tangle Evolution in Alzheimer's Disease," <i>Journal of Alzheimer's Disease</i> 2003, 5:65-77
	A30.	Hasegawa et al., "Protein Sequence and Mass Spectrometric Analysis of Tau in the Alzheimer's Disease Brain," <i>Journal of Biological Chemistry</i> 1992, 267(24):17047-17054
	A31.	Shutov et al., "[Diagnostic Significance of the type of In Vitro Interaction between Blood Lymphocytes and Serotonin in Multiple Sclerosis]" [Article in Russian], <i>Zh Nevrol Psikhiatr Im S S Korsakova</i> 2002, 102(4):35-38, Abstract only, from PubMed - PMID:12001663
	A32.	Lechin et al., "Plasma Neurotransmitters and Cortisol in Chronic Illness: Role of Stress," <i>J Medicine</i> 1994, 25(3-4):181-192, Abstract only, from PubMed - PMID:7996062
	A33.	Takahara et al., "Detection in Human Serum by Radioimmunoassay of Histidyl-Proline Diketopiperazine, a Metabolite of Thyrotropin-Releasing Hormone," <i>J Clinical Endocrinology</i> 1983, 56(2):312-319, Abstract only, from PubMed - PMID:6401750
	A34.	Prasad, "Bioactive Cyclic Dipeptides," <i>Peptides</i> 1995, 16:151-164
	A35.	Jicha et al., "Sequence Requirements for Formation of Conformational Variants of Tau Similar to Those Found in Alzheimer's Disease," <i>Journal of Neuroscience Research</i> 1999, 55:713-723
	A36.	Murray et al., "Role of α -Synuclein Carboxy-Terminus on Fibril Formation in Vitro," <i>Biochemistry</i> 2003, 42:8530-8540
	A37.	Steiner et al., "Histidyl Proline Diketopiperazine (Cyclo [His-Pro]) in Eating Disorders," <i>Neuropeptides</i> 1989, 14(3):185-189, Abstract only, from PubMed - PMID:2615922
	A38.	Prasad et al., "Isolation of cyclo(His-Pro)-like immunoreactivity from Human Urine and Demonstration of its Immunologic, Pharmacologic, and Physico-chemical Identity with the Synthetic Peptide," <i>Biochemistry Int</i> 1990, 21(3):425-434, Abstract only, from PubMed - PMID:2222490
	A39.	Hilton et al., "Food Contains the Bioactive Peptide, Cyclo(His-Pro), <i>J Clinical Endocrinol Metab</i> 1992, 75(2):375-378, Abstract only, from PubMed - PMID:1639938
↓	A40.	Banks et al., "Radioactively Iodinated Cyclo(His-Pro) Crosses the Blood-Brain Barrier and Reverses Ethanol-Induced Narcosis," <i>American J Physiol</i> 1993, 264(5 Pt 1):E723-729, Abstract only, from PubMed - PMID:8498494

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GE	A41.	Duntas et al., "A Fast Protein Liquid Chromatography (FPLC) Method for Study of Thyrotropin-releasing Hormone (TRH) and its metabolite Histidyl-Proline Diketopiperazine (CHP) in Human Blood: Degradation in Liver and Pancreatic Diseases," <i>Neuropeptides</i> 1993 25(6):357-361, Abstract only, from PubMed -PMID:8127415
	A42.	Shukla et al., "Role of Endogenous Cyclo(His-Pro) in Cold-Induced Hypothermia in the Desert Rat (<i>Mastomys natalensis</i>)," <i>Peptides</i> 1994, 15(8):1471-1474, Abstract only, from PubMed -PMID:7700849
	A43.	Jaspan et al., "Study of Passage of Peptides Across the Blood-Brain Barrier: Biological Effects of Cyclo(His-Pro) After Intravenous and Oral Administration, <i>Annals of the New York Academy of Science</i> 1994, 739:101-107, Abstract only, from PubMed -PMID:7832464
	A44.	Wolf et al., "Identification of Cyclo(His-Pro)-Like Immunoreactivity in Human Follicular Fluid: Correlation with Steroid and Peptide Hormones," <i>J Soc Gynecol Investigation</i> 1994, 1(3):220-224, Abstract only, from PubMed -PMID:9419775
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	A47.	Parker et al., "Evidence for the Presence of Immunoreactive Histidyl-Proline Diketopiperazine [Cyclo (His-Pro)] in the Adult Human Brain," <i>Peptides</i> 1983, 4(6):879-881, Abstract only, from PubMed -PMID:6672793
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	A49.	Lechan et al., "Thyrotropin Releasing Hormone but not Histidyl-Proline Diketopiperazine is Depleted from Rat Spinal Cord Following 5,7-Dihydroxytryptamine Treatment," <i>Brain Research</i> 1985, 326(1):152-155, Abstract only, from PubMed -PMID:3918765
	A50.	Diamanti Kandarakis et al., "Distribution and Characterization of Cyclo (His-Pro)-Like Immunoreactivity in the Human Gastrointestinal Tract," <i>Neuropeptides</i> 1985, 6(1):21-5, Abstract only, from PubMed -PMID:3990923
	A51.	Pekary et al., "In vitro Production of a TRH-Homologous Peptide and His-Pro Diketopiperazine by Human Semen," <i>J Androl</i> 1985, 6(6):379-385, Abstract only, from PubMed -PMID:3935636
	A52.	Koskinen, "Effect of Low Intravenous Doses of TRH, Acid-TRH and Cyclo (His-Pro) on Cerebral and Peripheral Blood Flows," <i>British Journal of Pharmacology</i> 1986, 87(3):509-519, Abstract only, from PubMed -PMID:3099875
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GE	A54.	Kurahashi et al., "[Histidyl-Proline Diketopiperazine (HPD) a Metabolite of Thyrotropin-Releasing Hormone (TRH), Improves the Ataxic Gait in 3-Acetylpyridine (3-AP) Treated Rats]" [Article in Japanese] <i>No To Shineki</i> 1986, 38(9):893-898, Abstract only, from PubMed -PMID:3790371
	A55.	Coggins et al., "High Affinity Specific Binding of the Thyrotrophin Releasing Hormone Metabolite Histidylproline to Rat Brain Membranes," <i>Neuropeptides</i> 1987, 9(1):83-91, Abstract only, from PubMed -PMID:3104816
	A56.	Mori et al., "Specific Radioimmunoassay of Cydo (His-Pro), a Biologically Active Metabolite of Thyrotropin-Releasing Hormone," <i>Endocrinology</i> 1981, 108(5):1995-1997, Abstract only, from PubMed -PMID:6783397
	A57.	Mori et al., "Regional Dissociation of Histidyl-Proline Diketopiperazine (Cydo-(His-Pro)) and Thyrotropin-Releasing Hormone (TRH) in the Rat Brain," <i>Brain Research</i> 1982, 231(2):451-453, Abstract only, from PubMed -PMID:6799149
	A58.	Prasad et al., "Distribution and Metabolism of Cyclo (His-Pro): a New Member of the Neuropeptide Family," <i>Peptides</i> 1982, 3(3):591-598, Abstract only, from PubMed -PMID:6812031
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	A61.	Gu et al., "Diketopiperazine Formation, Hydrolysis, and Epimerization of the New Dipeptide Angiotensin-Converting Enzyme Inhibitor RS-10085," <i>Pharm Res</i> 1987, 4(5):392-397, Abstract only, from PubMed -PMID:3508548
	A62.	Guerra et al., "PEGylation Prevents the N-Terminal Degradation of Megakaryocyte Growth and Development Factor," <i>Pharm Res</i> 1998, 15(12):1822-1827, Abstract only, from PubMed -PMID:9892464
	A63.	Sepetov et al., "Rearrangement, Racemization and Decomposition of Peptides in Aqueous Solution," <i>Peptide Research</i> 1991, 4(5):308-313, Abstract only, from PubMed -PMID:1802242
	A64.	Reubsaet et al., "Qualitative and Quantitative Aspects of the Degradation of Several Tripeptides Derived from the Antitumor Peptide Antagonist [Arg(6), D-Trp(7,9), MePhe(8)] Substance P[6-11]," <i>J Pharm Biomed Anal</i> 1999, 19(3-4):277-284, Abstract only, from PubMed -PMID:10704092
	A65.	Song et al., "Synergistic Antidiabetic Activities of Zinc, Cyclo (His-Pro), and Arachidonic Acid," <i>Metabolism</i> 2001 50(1):53-59, Abstract only, from PubMed -PMID:11172475
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GE	A67.	Pandey et al., "Synthetic Peptides Corresponding to a Repetitive Sequence of Malarial Histidine Rich Protein Bind Haem and Inhibit Haemozoin Formation in vitro," <i>Mol Biochem Parasitol</i> 1997, 90(1):281-287, Abstract only , from PubMed -PMID:9497049
	A68.	Baig et al., "High Performance Liquid Chromatography as a Tool in the Definition of Abnormalities in Monamine and Tryptophan Metabolites in Cerebrospinal Fluid from Patients with Neurological Disorders," <i>Biomed Chromatogr</i> 1991, 5(3):108-112, Abstract only , from PubMed -PMID:1863084
	A69.	Monaco et al., "Plasma and cerebrospinal fluid tryptophan in Multiple Sclerosis and Degenerative Diseases," <i>J Neurol Neurosurg Psychiatry</i> 1979 42(7):640-1, Abstract only , from PubMed -PMID:479903
	A70.	Scharpe et al., "Peptide Truncation by Dipeptidyl Peptidase IV: A New Pathway for Drug Discovery," <i>Verh K. Acad Geneesk Belg.</i> 2001, 63(1):5-32, Abstract only , from PubMed -PMID:11284388
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	A72.	Hilton et al., "Radioimmunoassay of Cyclo(His-Pro) in Unextracted Human Plasma: Report of a Normal Range and Definition of Factors Critical for Successful Assay," <i>Neuropeptides</i> 1989, 13(1):65-70, Abstract only , from PubMed -PMID:2922107
	A73.	Iriuchijima et al., "Thyrotropin-Releasing Hormone and Cyclo (His-Pro)-Like Immunoreactivities in the Cerebrospinal Fluids of 'Normal' Infants and Adults, and Patients with Various Neuropsychiatric and Neurologic Disorders," <i>Life Sci.</i> 1987, 41(22):2419-2428, Abstract only , from PubMed -PMID:2891013
	A74.	Hilton et al., "Relationship between Plasma Cyclo (His-Pro), a Neuropeptide Common to Processed Protein-Rich Food, C-Peptide/Insulin Molar Ratio in Obese Women," <i>Nutr Neurosci</i> 2001, 4(6):469-474, Abstract only , from PubMed -PMID:11843266
	A75.	Mori et al., "Brain TRH and Cyclo (His-Pro) and Brain Protein in the Newborn Rat are Altered by Maternal Liquid Protein Feeding," <i>Life Sci</i> 1983, 32(14):1607-1612, Abstract only , from PubMed -PMID:6403790
	A76.	Mori et al., "[TRH and Cyclo (His-Pro) Concentrations in the Young Rat Brain are Altered by a Liquid Protein Diet]" [Article in Japanese], <i>Nippon Naibunpi Gakkai Zasshi</i> 1987, 63(7):846-852
	A77.	Mori et al., "Alteration by Liquid Protein Diet of TRH and Cyclo(His-Pro) in the Young Rat Brain," <i>Res. Commun Chem Pathol Pharmacol</i> 1985, 47(1):157-160, Abstract only , from PubMed -PMID:392073
↓	A78.	Goolcharan et al., "Comparison of the Rates of Deamidation, Diketopiperazine Formation and Oxidation in Recombinant Human Vascular Endothelial Growth Factor and Model Peptides," <i>AAPS PharmSci</i> 2000, 2(1)E5, Abstract only , from PubMed - PMID:11741221

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	A80.	Hilton et al., "Identification and Characterization of Cyclo (His-Pro)-Like Immunoreactivity in Amniotic Fluid," <i>Peptides</i> 10(2):299-301, Abstract only , from PubMed - PMID:2755872
	A81.	Bhargava et al., "Inhibition of Neuroleptic-Induced Dopamine Receptor Supersensitivity by Cyclo (Leu-Gly)," <i>Pharmacol Biochem Behav</i> 1980, 13(5):633-636, Abstract only , from PubMed - PMID:7443732
	A82.	Ledduque et al., "Histidyl-Proline Diketopiperazine (His-Pro DKP) Immunoreactivity is Present in the Glucagon-Containing Cells of the Human Fetal Pancreas," <i>J Clin Invest</i> 1987, 79(3):875-880, Abstract only , from PubMed - PMID:3102558
	A83.	Battersby et al., "Diketopiperazine Formation and N-Terminal Degradation in Recombinant Human Growth Hormone," <i>Int J Peptide Protein Res</i> 1994, 44(3):215-222, Abstract only , from PubMed - PMID:7822097
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	A85.	Yanagisawa et al., "The Subcellular and Organ Distribution and Natural Form of Histidyl-Proline Diketopiperazine in Rat brain Determined by a Specific Radioimmunoassay," <i>J Biol Chem</i> 1980, 255(21):10290-10294, Abstract only , from PubMed - PMID:7430126
	A86.	Hoffman et al., "An Enzymatically Stable Peptide with Activity in the Central Nervous System: Its Penetration Through to Blood-CSF Barrier," <i>Brain Res.</i> 1977, 122(1):87-94, Abstract only , from PubMed - PMID:837226
	A87.	Meester et al., "In Vivo Inhibition of Dipeptidyl Peptidase IV Activity by Pro-Pro-diphenyl-phosphonate (Prodipine)," <i>Biochemical Pharmacology</i> 1997, 54:173-179
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	A89.	Wilber et al., "Histidyl-proline diketopiperazine: a potent and chronic appetite-inhibiting neuropeptide," <i>Trans Assoc. Am Physicians</i> 1986, 99:245-249
	A90.	Wilber et al., "Endogenous histidyl-proline diketopiperazine [cyclo (His-Pro)]: a potential satiety neuropeptide in normal and genetically obese rodents," <i>Trans Assoc Am Physicians</i> 1983, 96:131-136
	A91.	Bhargava, "Inhibition of abstinence syndrome in opiate dependent mice by cyclo (His-Pro)," <i>Life Sci</i> 1981, 28(11):1261-1267
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GE	A93.	Bhargava, "Antagonism of ketamine-induced anesthesia and hypothermia by thyrotropin releasing hormone and cyclo (His-Pro)," <i>Neuropharmacology</i> 1981, 20(7):699-702
	A94.	Mori et al., "Histidyl-Proline Diketopiperazine cyclo (His-Pro): measurement by radioimmunoassay in human blood in normal subject and in patients with hyper- and hypothyroidism," <i>Biochem Biophys Res Commun</i> 1982, 109(2):541-547
	A95.	Luca et al., "Determination of serotonin content and ceruloplasmin activity, of blood and CSF amino acid level in multiple sclerosis," <i>Neurol Psychiatr (Bucur)</i> 1986, 24(3):153-159
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				Art Unit	1649
				Examiner Name	G. S. Emch
				Attorney Docket Number	4172-85

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Examiner Initials*	Cite No. ¹	Document Number Number-kind Code ² (if known)	Publication Date MM-DD-YYYY	Name of Patentee of Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
GE		US-6,815,214 B2	11-09-2004	Boyce, et al.	
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		PCT -				
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OTHER ART (Including Author, Title, Date, Pertinent Pages, etc.)

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↓		Wennemers et al., Diketopiperazine Receptors: A Novel Class of Highly Selective Receptors for Binding Small Peptides, Chem. Eur. J. 2001, Vol. 7, No. 15, pages 3342-3347.
↓		Prakash et al., Synthesis and Biological Activity of Novel Neuroprotective Diketopiperazines, Bioorganic & Medicinal Chemistry, September 2002, Vol. 10, No. 9, pages 3043-3048.
↓		McClelland et al., An investigation into the biological activity of the selected histidine-containing diketopiperazines cyclo(His-Phe) and cyclo(His-Tyr), Journal of Pharmacy and Pharmacology, September 2004, Vol. 56, No. 9, pages 1143-1153.

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Filing Date	2003-10-02
First Named Inventor	David Bar-Or
Art Unit	1649
Examiner Name	Gregory Emch
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